

**INFORMATION DISCLOSURE
STATEMENT BY APPLICANT**
(Not for submission under 37 CFR 1.99)

Application Number		10521040
Filing Date		2005-06-16
First Named Inventor		Herman Jan Tijmen Coelingh Bennink et al.
Art Unit		1616
Examiner Name		Mei-Ping Chui
Attorney Docket Number		0470-050079

U.S. PATENTS						<input type="button" value="Remove"/>
Examiner Initial*	Cite No	Patent Number	Kind Code†	Issue Date	Name of Patentee or Applicant of cited Document	Pages, Columns, Lines where Relevant Passages or Relevant Figures Appear
1	6500814			2002-12-31	Hesch	
2	6214815			2001-04-10	Shangold et al.	
3	5827843			1998-10-27	Koninckx	
4	5662927			1997-09-02	Ehrlich et al.	
5	5633242			1997-05-27	Oettel et al.	
6	5340586			1994-06-23	Pike et al.	
7	5223261			1993-06-29	Nelson et al.	
8	5130137			1992-07-14	Crowley, Jr.	

**INFORMATION DISCLOSURE
STATEMENT BY APPLICANT**
(Not for submission under 37 CFR 1.99)

Application Number	10521040
Filing Date	2005-06-16
First Named Inventor	Herman Jan Tijmen Coelingh Bennink et al.
Art Unit	1616
Examiner Name	Mei-Ping Chui
Attorney Docket Number	0470-050079

9	5063507		1991-11-05	Lindsey et al.	
10	4762717		1988-06-09	Crowley, Jr.	
11	4722941		1988-02-02	Eckert et al.	
12	4624665		1986-11-25	Nuwayser	
13	4573996		1986-03-04	Kwiatek et al.	
14	4460372		1984-07-17	Campbell et al.	
15	3797494		1974-03-19	Zaffaroni	
16	3440320		1969-04-22	Sackler et al.	

If you wish to add additional U.S. Patent citation information please click the Add button.

U.S. PATENT APPLICATION PUBLICATIONS

Examiner Initial*	Cite No	Publication Number	Kind Code ¹	Publication Date	Name of Patentee or Applicant of cited Document	Pages, Columns, Lines where Relevant Passages or Relevant Figures Appear
1		20020183299		2002-12-05	Voskuhl	

If you wish to add additional U.S. Published Application citation information please click the Add button.

**INFORMATION DISCLOSURE
STATEMENT BY APPLICANT**
(Not for submission under 37 CFR 1.99)

Application Number	10521040
Filing Date	2005-06-16
First Named Inventor	Herman Jan Tijmen Coelingh Bennink et al
Art Unit	1616
Examiner Name	Mei-Ping Chui
Attorney Docket Number	0470-050079

FOREIGN PATENT DOCUMENTS							Remove	
Examiner Initial*	Cite No	Foreign Document Number ³	Country Code ² ^j	Kind Code ⁴	Publication Date	Name of Patentee or Applicant of cited Document	Pages, Columns, Lines where Relevant Passages or Relevant Figures Appear	T ⁵
1	0185154	WO	A2	2001-11-15	Oregon Health Sciences University et al.			<input type="checkbox"/>
2	1700602	EP	A1	2001-05-18	Pantarhei Bioscience B.V			<input type="checkbox"/>
3	0130357	WO	A	2001-05-03	Theramex			<input type="checkbox"/>
4	0073416	WO	A1	2000-12-07	University of Cincinnati			<input type="checkbox"/>
5	0062753	WO		2000-10-26	Jenapharm GmbH & Co.			<input type="checkbox"/>
6	19917930	DE	A1	2000-10-19	Schering			<input type="checkbox"/>
7	9858657	WO	A1	1998-12-30	Mannaerts et al			<input type="checkbox"/>
8	9603929	WO	A1	1996-02-15	Blex, Inc.			<input type="checkbox"/>
9	9517895	WO		1995-07-06	Ehrlich et al			<input type="checkbox"/>

**INFORMATION DISCLOSURE
STATEMENT BY APPLICANT**
(Not for submission under 37 CFR 1.99)

Application Number	10521040
Filing Date	2005-06-16
First Named Inventor	Herman Jan Tijmen Coelingh Bennink et al.
Art Unit	1616
Examiner Name	Mei-Ping Chui
Attorney Docket Number	0470-050079

10	9502408	WO	A1	1995-01-26	Schering	<input type="checkbox"/>
11	9218107	WO	A1	1992-10-29	University of Southern California	<input type="checkbox"/>
12	468690	EP	A1	1991-07-17	Sumitomo Chemical Co., Ltd.	<input type="checkbox"/>
13	0402950	EP	A	1975-12-18	Orion Yhtymae OY	<input type="checkbox"/>
14	2426779	DE	A1	1975-12-18	Schering AG	<input type="checkbox"/>
15	2336434	DE	A1	1975-04-17	Schering AG	<input type="checkbox"/>
16	2336433	DE	A1	1975-04-03	Schering AG	<input type="checkbox"/>
17	9603929	WO	A1	1966-02-15	Biox, Inc.	<input type="checkbox"/>

If you wish to add additional Foreign Patent Document citation information please click the Add button

Add

NON-PATENT LITERATURE DOCUMENTS

Remove

Examiner Initials*	Cite No	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc), date, pages(s), volume-issue number(s), publisher, city and/or country where published.	T6
	1	VISSEER et al., "In vitro effects of estrelol on receptor binding, drug targets and human liver cell metabolism," CLIMACTERIC (2008) 11(1) Appx. II: 1-5.	<input type="checkbox"/>

**INFORMATION DISCLOSURE
STATEMENT BY APPLICANT**
(Not for submission under 37 CFR 1.99)

Application Number	10521040
Filing Date	2005-06-16
First Named Inventor	Herman Jan Tijmen Coelingh Bennink et al.
Art Unit	1616
Examiner Name	Mei-Ping Chui
Attorney Docket Number	0470-050079

2	VISSEER et al., "First human exposure to exogenous single-dose oral estetrol in early postmenopausal women," CLIMACTERIC (2008) 11(1): 1-10.	<input type="checkbox"/>
3	VISSEER et al., "Clinical applications of estetrol," J. Of Steroid Biochem and Molecular Biol. (2009) 114: 85-89.	<input type="checkbox"/>
4	HOLINKA et al., "Estetrol: A unique steroid in human pregnancy," J. of Steroid Biochem and Molecular Biol. (2009) 110: 138-143.	<input type="checkbox"/>
5	COELINGH BENNINK et al., "Oral bioavailability and bone sparing effects of estetrol in an osteoporosis model," CLIMACTERIC (2008) 11 (Supp 3): 1-13.	<input type="checkbox"/>
6	ALBERTAZZI PAOLA et al., "The Effect of Tibolone Versus Continuous Combined Norethisterone Acetate and Oestradiol on Memory, Libido and Mood of Postmenopausal Women: A Pilot study"; Database Biosis 'Online'; 10/31/2000; pages 223-229; Vol. 36, No. 3; Biosciences Information Service, Philadelphia, PA., U.S.	<input type="checkbox"/>
7	ALLEN ET AL., An Ovarian Hormone: Preliminary Report on Its Localization, Extraction and Partial Purification, and Action in Test Animals, JAMA, September 8, 1923, Vol. 81, pp. 819-821.	<input type="checkbox"/>
8	ALLEN ET AL., The Induction of a Sexually Mature Condition in Immature Females by Injection of the Ovarian Follicular Hormone, Am. J. Physiol., 1924, Vol. 69, pp. 577-588.	<input type="checkbox"/>
9	JONES ET AL., The Effects of Various Steroids on the Vaginal Histology in the Rat, Fertility and Sterility, April 1973, Vol. 24, No. 4, pp. 284-291.	<input type="checkbox"/>
10	TULCHINSKY ET AL., Plasma Esterol as an Index of Fetal Well-Being, J. Clin. Endocrinol. Metab., 1975, Vol 40. Pp. 560-567	<input type="checkbox"/>
11	HAMMOND ET AL., A Versatile Method for the Determination of Serum Cortisol Binding Globulin and Sex Hormone Binding Globulin Binding Capacities, 1983, Vol. 132, pp. 101-110.	<input type="checkbox"/>
12	ELGER ET AL., Sulfamates of Various Estrogens are Prodrugs with Increased Systemic and Reduced Hepatic Estrogenicity at Oral Application, J. Steroid Biochem. Molec. Biol., 1995, Vol. 55, No. 3 / 4, pp. 395-403.	<input type="checkbox"/>

**INFORMATION DISCLOSURE
STATEMENT BY APPLICANT**
(Not for submission under 37 CFR 1.99)

Application Number	10521040
Filing Date	2005-06-16
First Named Inventor	Herman Jan Tijmen Coelingh Bennink et al
Art Unit	1616
Examiner Name	Mei-Ping Chui
Attorney Docket Number	0470-050079

13	MURPHY ET AL., Endometrial Effects of Long-Term Low-Dose Administration of RU486, Fertility and Sterility, April 1995, Vol. 63, No. 4, pp. 761-766.	<input type="checkbox"/>
14	REEL ET AL., Survey and Assessment of Mammalian Estrogen Biological Assays for Hazard Characterization, Fundamental and Applied Toxicology, 1996, Vol. 34, pp. 288-305.	<input type="checkbox"/>
15	BERAL ET AL., Use of HRT and the Subsequent Risk of Cancer, Journal of Epidemiology and Biostatistics, 1999, Vol. 4, No. 3, pp. 191-215.	<input type="checkbox"/>
16	TAVANI ET AL., The Adverse Effects of Hormone Replacement Therapy, Drugs & Aging, May 1999, Vol. 14, No. 5, pp. 347-357.	<input type="checkbox"/>
17	PIKE ET AL., Progestins and Menopause: Epidemiological Studies of Risks of Endometrial and Breast Cancer, Steroids, 2000, Vol. 65, pp. 359-664.	<input type="checkbox"/>
18	AVVAKUMOV ET AL., Steroid-binding Specificity of Human Sex Hormon-binding Globulin Is Influenced by Occupancy of a Zinc-binding Site, The Journal of Biological Chemistry, August 25, 2000, Vol. 275, No. 34, pp. 25920-25925.	<input type="checkbox"/>
19	HOLINKA, ET AL., "Comparison of Effects of Esterol and Tamoxifen with Those of Estradiol and Estradiol on the Immature Rat Uterus", Biology of Reproduction, Society for the Reproduction Society for the Study of Reproduction, Champaign, IL, US, 1980, Vol. 22, No. 4, pp. 913-926.	<input type="checkbox"/>
20	JANSSON ET AL., "Estrogen Induces a Potent Suppression of Experimental Autoimmune Encephalomyelitis and Collagen-Induced Arthritis in Mice", Journal of Neuroimmunology, Elsevier Science Publishers BV XX, 1994, Vol. 53, No. 2, pp. 203-207.	<input type="checkbox"/>
21	ERDBRUEGGER ET AL., Drug Discovery Today: Disease Mechanisms (2004), Vol. 1, pp. 73-81.	<input type="checkbox"/>
22	Lab Tests Online (www.labtestsonline.org/understanding/conditions/autoimmune.html) retrieved on 10/15/2009.	<input type="checkbox"/>
23	www.tiscali.co.uk/lifestyle/healthfitness/health_advice/meddoctor/archive/000489.html , 10/15/2009.	<input type="checkbox"/>

**INFORMATION DISCLOSURE
STATEMENT BY APPLICANT**
(Not for submission under 37 CFR 1.99)

Application Number	10521040
Filing Date	2005-06-16
First Named Inventor	Herman Jan Tijmen Coelingh Bennink et al.
Art Unit	1616
Examiner Name	Mei-Ping Chui
Attorney Docket Number	0470-050079

24	MedlinePlus Medical Encyclopedia: Multiple Sclerosis, retrieved on 03/28/2008 via www.nlm.nih.gov/medlineplus/ency/article/000737.htm , dated on 08/06/2007, Page 1 and 2; also see WebMD: Multiple Sclerosis - Prevention, retrieved on 03/28/2008 via www.webmd.com/multiple-sclerosis/tc/multiple-sclerosis-ms-prevention , dated on 03/23/2006.	<input type="checkbox"/>
25	MedlinePlus Medical Encyclopedia: rheumatoid arthritis, retrieved on 03/28/2008 via www.nlm.nih.gov/medlineplus/ency/article/000431.htm , dated on 07/27/2007, page 1-2 and 4; also WebMD: Rheumatoid Arthritis - Prevention, retrieved on 03/28/2008 via www.webmd.com/rheumatoid-arthritis/tc/rheumatoid-arthritis-prevention , dated on 08/23/2006.	<input type="checkbox"/>
26	MUECK et al., "Angio and Anti-Angiogenetic Effects of Estradiol and its Metabolites", <i>J. Clin. Basic Cardiol.</i> , 2001, pp. 153-155, Vol. 4, No. 2.	<input type="checkbox"/>
27	SHAH et al., "Estrogen and Skin. An Overview", <i>Am. J. Clin. Dermatol.</i> , 2001, pp. 143-150, Vol. 2, No. 3.	<input type="checkbox"/>
28	SITRUK-WARE et al., "Local Hormonal Treatment for Urogenital Atrophy After Menopause", <i>Shweiz. Rundsch. Med. Praxis</i> , 1997, pp. 1245-1248, Vol. 86, No. 33, and SITRUK-WARE, English Translation, 1997. <i>Praxis, Schweizerische Rundschau für Medizin</i> , Vol. 86, No. 33, pp 1-13	<input type="checkbox"/>
29	SCHMIDT et al., "Treatment of Skin Aging with Topical Estrogens", <i>Int. J. Dermatol.</i> , 1996, pp. 669-674, Vol. 35, No. 9.	<input type="checkbox"/>
30	YOUNGLAI ET AL., <i>Journal of Clinical Endocrinology and Metabolism</i> , 1968, Vol. 28, Issue 11, pp 1611-1617.	<input type="checkbox"/>
31	Webster Ninth New Collegiate Dictionary, 2000, Definition of Prevention, pg 1.	<input type="checkbox"/>
32	WILLHITE ET AL., <i>Pharmacotherapy</i> , 2001, Vol. 21, Issue 4, pp 464-480.	<input type="checkbox"/>
33	SEEGER et al., "The inhibitory effect of endogenous estrogen metabolites on copper-mediated in vitro oxidation of LDL", <i>Int. Journal of Clinical Pharmacology and Therapeutics</i> , (1998), Vol. 36, No. 7, pp 383-385.	<input type="checkbox"/>
34	KUIPERS ET AL., "Enterohepatic Circulation in the Rat", <i>Gastroenterol.</i> , Vol. 88, pp 403-411 (1985).	<input type="checkbox"/>

**INFORMATION DISCLOSURE
STATEMENT BY APPLICANT**
(Not for submission under 37 CFR 1.99)

Application Number	10521040
Filing Date	2005-06-16
First Named Inventor	Herman Jan Tijmen Coelingh Bennink et al.
Art Unit	1616
Examiner Name	Mei-Ping Chui
Attorney Docket Number	0470-050079

	35	SCHWARTZ, "A Model for the Regulation of Ovulation in the Rat", Recent Prog. Horm. Res., Vol. 25, pp 1-55, (1969). <input type="checkbox"/>
	36	BEATTIE ET AL., "The Differential Effects of Diestrous Progestogen Administration on Proestrous Gonadotrophin Levels, Endocrinol", Vol 97, pp. 885-890, (1975). <input type="checkbox"/>
	37	DE VISSER ET AL., "Endocrinological Studies with (7a, 17 a)-17-Hydroxy-7-me norpregn-5(10)-en-20-yn-3-one (Org OD 14), Arzneim, Forsh., Vol. 34, pp 1010-1020, (1984). <input type="checkbox"/>

If you wish to add additional non-patent literature document citation information please click the Add button

EXAMINER SIGNATURE

Examiner Signature	Date Considered
--------------------	-----------------

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through a citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹ See Kind Codes of USPTO Patent Documents at www.USPTO.GOV or MPEP 901.04. ² Enter office that issued the document, by the two-letter code (WIPO Standard ST.3). ³ For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. ⁴ Kind of document by the appropriate symbol as indicated on the document under WIPO Standard ST.16 if possible. ⁵ Applicant is to place a check mark here if English language translation is attached.